PATENT

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re application of: BRAJ BHUSHAN **LOHRAY,** et al

For: NOVEL BYCYCLIC COMPOUNDS AND THEIR USE IN MEDICINE; PROCESS FOR THEIR PREPARATION AND PHARMACEUTICAL COMPOSITIONS C CONTAINING THEM

Attorney Docket No.: U 013323-5

Commissioner of Patents and Trademarks
Washington, D.C. 20231

Sir:

PRELIMINARY AMENDMENT

IN THE CLAIMS

Please cancel Claims 1 – 28.

Please add claims 29 – 31.

CERTIFICATE UNDER 37 CFR 1.10

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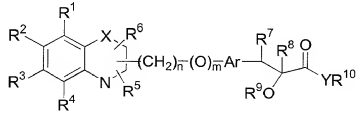
-- 29. A method for preventing or treating complications of diabetes said diabetes caused by insulin resistance or impaired glucose tolerance, said method comprising administering a therapeutically effective amount of a compound of formula (I)



(I)

its tautomeric forms, its stereoisomers, its polymorphs, its pharmaceutically acceptable salts, or its pharmaceutically acceptable solvates, wherein the groups R^1, R^2, R^3, R^4, and the groups R^5 and R^6 when attached to a carbon atom, may be same or different and represent hydrogen, halogen, hydroxy, nitro, cyano, formyl or optionally substituted groups selected from alkyl, cycloalkyl, alkoxy, cycloalkoxy, aryl, aryloxy, aralkyl, aralkoxy, heterocyclyl selected from aziridinyl, pyrrolidinyl, morpholinyl, piperidinyl or piperazinyl; heteroaryl group selected from pyridyl, thienyl, furyl, pyrrolyl, oxazolyl, thiazolyl, imidazolyl, oxadiazolyl, tetrazolyl, benzopyranyl, or benzofuranyl; heteroaralkyl group selected from furanmethyl, pyridinemethyl, oxazolemethyl, or oxazolethyl; heteroaryloxy wherein the heteroaryl moiety is as defined above, heteroaralkoxy; wherein the heteroaralkyl moiety is as defined above; acyl selected from acetyl, propionyl or benzoyl; acyloxy, hydroxyalkyl, amino, acylamino, alkylamino, arylamino, aralkylamino, aminoalkyl, alkoxycarbonyl, aryloxycarbonyl, aralkoxycarbonyl, alkoxyalkyl, aryloxyalkyl, aralkoxyalkyl, alkylthio, thioalkyl, alkoxycarbonylamino, aryloxycarbonylamino, aralkoxycarbonylamino, carboxylic acid or its esters or amides, or sulfonic acid or its esters or amides; one or both of R^5 and R^6 may represent an oxo group when attached to a carbon atom; R^5 and R^6 when attached to a nitrogen atom may be the same or different and represent hydrogen, hydroxy, formyl or

optionally substituted groups selected from alkyl, cycloalkyl, alkoxy, cycloalkoxy, aryl, aralkyl, heterocyclyl selected from aziridinyl, pyrrolidinyl, morpholinyl, piperidinyl or piperazinyl; heteroaryl group selected from pyridyl, thienyl, furyl, pyrrolyl, oxazolyl, thiazolyl, imidazolyl, oxadiazolyl, tetrazolyl, benzopyranyl, or benzofuranyl; heteroaralkyl group selected from furanmethyl, pyridinemethyl, oxazolemethyl, or oxazolethyl; acyl selected from acetyl, propionyl or benzoyl; acyloxy, hydroxyalkyl, amino, acylamino, alkylamino, arylamino, aralkylamino, aminoalkyl, aryloxy, aralkoxy, heteroaryloxy wherein the heteroaryl moiety is as defined above; heteroaralkoxy, wherein the heteroaralkyl moiety is as defined above; alkoxycarbonyl, aryloxycarbonyl, aralkoxycarbonyl, alkoxyalkyl, aryloxyalkyl, aralkoxyalkyl, alkylthio, thioalkyl groups, esters or amides of carboxylic acid or esters or amides of sulfonic acid; X represents oxygen or sulfur; Ar represents an optionally substituted divalent single aromatic or fused aromatic or heterocyclic group; R^7 represents hydrogen atom, hydroxy, alkoxy, halogen, lower alkyl, optionally substituted aralkyl group or forms a bond together with R^8; R^8 represents hydrogen, hydroxy, alkoxy, halogen, lower alkyl group, acyl, or optionally substituted aralkyl or R^8 forms a bond together with R^7; R^9 represents hydrogen, or optionally substituted groups selected from alkyl, cycloalkyl, aryl, aralkyl, alkoxyalkyl, alkoxycarbonyl, aryloxycarbonyl, alkylaminocarbonyl, arylaminocarbonyl, acyl selected from acetyl, propionyl or benzoyl; heterocyclyl selected from aziridinyl, pyrrolidinyl, morpholinyl, piperidinyl or piperazinyl; heteroaryl group selected from pyridyl, thienyl, furyl, pyrrolyl, oxazolyl, thiazolyl, imidazolyl, oxadiazolyl, tetrazolyl, benzopyranyl, or benzofuranyl; or heteroaralkyl group selected from furanmethyl, pyridinemethyl, oxazolemethyl, or oxazolethyl; R^{10} represents hydrogen or optionally substituted groups selected from alkyl, cycloalkyl, aryl, aralkyl, heterocyclyl selected from aziridinyl, pyrrolidinyl, morpholinyl, piperidinyl or piperazinyl; heteroaryl group selected

from pyridyl, thienyl, furyl, pyrrolyl, oxazolyl, thiazolyl, imidazolyl, oxadiazolyl, tetrazolyl, benzopyranyl, or benzofuranyl; or heteroaralkyl group selected from furanmethyl, pyridinemethyl, oxazolemethyl or oxazolethyl; Y represents oxygen or NR^{12}, where R^{12} represents hydrogen, alkyl, aryl, hydroxyalkyl, aralkyl, heterocyclyl selected from aziridinyl, pyrrolidinyl, morpholinyl, piperidinyl or piperazinyl; heteroaryl group selected from pyridyl, thienyl, furyl, pyrrolyl, oxazolyl, thiazolyl, imidazolyl, oxadiazolyl, tetrazolyl, benzopyranyl, or benzofuranyl; or heteroaralkyl group selected from furanmethyl, pyridinemethyl, oxazolemethyl or oxazolethyl; R^{10} and R^{12} together may form a 5 or 6 membered cyclic structure containing carbon atoms, which may optionally contain one or more heteroatoms selected from oxygen, sulfur or nitrogen; the linking group represented by $-(CH_2)_n-(O)_m-$ may be attached either through a nitrogen atom or a carbon atom; n is an integer ranging from 1-4 and m is an integer 0 or 1, with the proviso that when the linking group is attached through a carbon atom and either of R^5 or R^6 represents an oxo group and Y is an oxygen atom, R^9 does not represent a hydrogen atom; and a pharmaceutically acceptable carrier, diluent or excipient to a patient in need thereof, wherein the complication is hyperglycemia, osteoporis, hyperlidemia, nephrotic syndrome or disorders related to endothelial cell division.

-- 30. A method for preventing or treating complications of diabetes said diabetes caused by insulin resistance or impaired glucose tolerance said method comprising administering a therapeutically effective amount of a compound selected from:

Ethyl (E/Z)-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-ethoxypropenoate;

(±) Methyl 3-[4-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]

-2-ethoxypropanoate;

(+) Methyl 3-[4-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]
-2-ethoxypropanoate;

(-) Methyl 3-[4-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]
-2-ethoxypropanoate;

Ethyl (E/Z)-3-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)methylbenzofuran-5-yl]
2-ethoxypropenoate;

Ethyl (E/Z)-3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-
ethoxypropenoate;

(±) Methyl 3-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)methylbenzofuran-5-yl]
-2-ethoxypropanoate;

(+) Methyl 3-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)methylbenzofuran-5-yl]
-2-ethoxypropanoate;

(-) Methyl 3-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)methylbenzofuran-5-yl]
-2-ethoxypropanoate;

(±) Methyl-3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-
ethoxypropanoate;

(+) Methyl-3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-
ethoxypropanoate;

(-) Methyl-3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-ethoxypropanoate;

(±) Methyl 2-methyl-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy] phenyl]-2-ethoxypropanoate;

(+) Methyl 2-methyl-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy] phenyl]-2-ethoxypropanoate;

(-) Methyl 2-methyl-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy] phenyl]-2-ethoxypropanoate;

(±) Methyl 2-(2-fluorobenzyl)-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-ethoxypropanoate;

(+) Methyl 2-(2-fluorobenzyl)-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-ethoxypropanoate;

(-) Methyl 2-(2-fluorobenzyl)-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-ethoxypropanoate;

Ethyl (E/Z)-3-[4-[2-(3-oxo-2H-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-ethoxypropenoate;

(±) Methyl 3-[4-[2-(3-oxo-2H-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-ethoxypropanoate;

(+) Methyl 3-[4-[2-(3-oxo-2H-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-ethoxypropanoate;

(-) Methyl 3-[4-[2-(3-oxo-2H-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-ethoxypropanoate;

Ethyl (E/Z)-3-[6-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]naphthyl]-2-ethoxypropenoate;

(±) Methyl 3-[6-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]naphthyl]-2-ethoxypropanoate;

(+) Methyl 3-[6-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]naphthyl]-2-ethoxypropanoate;

(-) Methyl 3-[6-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]naphthyl]-2-ethoxypropanoate;

(±) Ethyl 3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-hydroxypropanoate;

(+)Ethyl 3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-hydroxypropanoate;

(-)Ethyl 3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-hydroxypropanoate;

(±) Ethyl 3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-hydroxypropanoate;

(+)Ethyl 3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-hydroxypropanoate;

(-) Ethyl 3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-hydroxypropanoate;

(±) Ethyl 3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-benzyloxypropanoate;

(+) Ethyl 3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-benzyloxypropanoate;

(-)Ethyl 3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-benzyloxypropanoate;

(±) Ethyl 3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-butoxypropanoate;

(+) Ethyl 3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-butoxypropanoate;

(-)Ethyl 3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-butoxypropanoate;

(±) Ethyl 3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-hexyloxy propanoate;

(+) Ethyl 3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-hexyloxy propanoate;

(-) Ethyl 3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-hexyloxy propanoate;

Ethyl (E/Z)-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-phenoxypropenoate;

(±) Methyl 3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-phenoxypropanoate;

(+) Methyl 3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-phenoxypropanoate;

(-) Methyl 3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-phenoxypropanoate;

Ethyl (E/Z)-3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-phenoxypropenoate;

(±) Methyl 3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-phenoxypropanoate;

(+) Methyl 3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-phenoxypropanoate;

(-) Methyl 3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-phenoxypropanoate;

Ethyl (E/Z)-3-[4-(4-methyl-3,4-dihydro-2H-1,4-benzoxazin-2-yl) methoxyphenyl] -2-ethoxypropenoate;

(±) Methyl 3-[4-(4-methyl-3,4-dihydro-2H-1,4-benzoxazin-2-yl) methoxyphenyl] -2-ethoxypropanoate;

(+) Methyl 3-[4-(4-methyl-3,4-dihydro-2H-1,4-benzoxazin-2-yl) methoxyphenyl]

-2-ethoxypropanoate;

(-) Methyl 3-[4-(4-methyl-3,4-dihydro-2H-1,4-benzoxazin-2-yl) methoxyphenyl] -2-ethoxypropanoate;

Ethyl (E/Z)-3-[4-(4-benzyl-3,4-dihydro-2H-1,4-benzoxazin-2-yl) methoxyphenyl] -2-ethoxypropenoate;

(±) Methyl 3-[4-(4-benzyl-3,4-dihydro-2H-1,4-benzoxazin-2-yl) methoxyphenyl] -2-ethoxypropanoate;

(+) Methyl 3-[4-(4-benzyl-3,4-dihydro-2H-1,4-benzoxazin-2-yl) methoxyphenyl] -2-ethoxypropanoate;

(-) Methyl 3-[4-(4-benzyl-3,4-dihydro-2H-1,4-benzoxazin-2-yl) methoxyphenyl] -2-ethoxypropanoate;

(±) 3-[4-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-ethoxypropanoic acid and its salts;

(+) 3-[4-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-ethoxypropanoic acid and its salts;

(-) 3-[4-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-ethoxypropanoic acid and its salts;

(±) 3-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)methylbenzofuran-5-yl]-2-ethoxypropanoic acid and its salts;

(+) 3-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)methylbenzofuran-5-yl]-2-ethoxypropanoic acid and its salts;

(-) 3-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)methylbenzofuran-5-yl]-2-ethoxypropanoic acid and its salts;

(±) 3-[2-(2,3-Dihydro-1,4-benzothiazin-4-yl)methylbenzofuran-5-yl]-2-ethoxypropanoic acid and its salts;

(+) 3-[2-(2,3-Dihydro-1,4-benzothiazin-4-yl)methylbenzofuran-5-yl]-2-ethoxypropanoic acid and its salts;

(-) 3-[2-(2,3-Dihydro-1,4-benzothiazin-4-yl)methylbenzofuran-5-yl]-2-ethoxypropanoic acid and its salts;

(±) 3-[4-[2-(2,3-Dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-ethoxypropanoic acid and its salts;

(+) 3-[4-[2-(2,3-Dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-ethoxypropanoic acid and its salts;

(-) 3-[4-[2-(2,3-Dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-ethoxypropanoic acid and its salts;

(±) 3-[4-[2-(2,3-Dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-ethoxypropanamide;

(+) 3-[4-[2-(2,3-Dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-ethoxypropanamide;

(-) 3-[4-[2-(2,3-Dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-

ethoxypropanamide;

(±) N-Methyl-3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl] -2-ethoxypropanamide;

(+) N-Methyl-3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl] -2-ethoxypropanamide;

(-) N-Methyl-3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl] -2-ethoxypropanamide;

(±) 3-[4-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-ethoxypropanamide;

(+) 3-[4-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-ethoxypropanamide;

(-) 3-[4-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-ethoxypropanamide;

(±) N-Methyl-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-ethoxypropanamide;

(+) N-Methyl-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-ethoxypropanamide;

(-) N-Methyl-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-ethoxypropanamide;

(±) N-Benzyl-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]

-2-ethoxypropanamide;

(+) N-Benzyl-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]

-2-ethoxypropanamide;

(-) N-Benzyl-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]

-2-ethoxypropanamide;

(±) N-Benzyl-3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]

-2-ethoxypropanamide;

(+) N-Benzyl-3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]

-2-ethoxypropanamide;

(-) N-Benzyl-3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]

-2-ethoxypropanamide;

(±) 2-Methyl-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]

-2-ethoxypropanoic acid and its salts;

(+) 2-Methyl-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]

-2-ethoxypropanoic acid and its salts;

(-) 2-Methyl-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]

-2-ethoxypropanoic acid and its salts;

(±) 2-(2-Fluorobenzyl)-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]

phenyl]-2-ethoxypropanoic acid and its salts;

(+) 2-(2-Fluorobenzyl)-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]

phenyl]-2-ethoxypropanoic acid and its salts;

(-) 2-(2-Fluorobenzyl)-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy] phenyl]-2-ethoxypropanoic acid and its salts;

(±) 3-[4-[2-(3-Oxo-2H-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-ethoxypropanoic acid and its salts;

(+) 3-[4-[2-(3-Oxo-2H-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-ethoxypropanoic acid and its salts;

(-) 3-[4-[2-(3-Oxo-2H-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-ethoxypropanoic acid and its salts;

(±) 3-[4-[2-(3-Oxo-2H-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-ethoxypropanoic acid and its salts;

(+) 3-[4-[2-(3-Oxo-2H-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-ethoxypropanoic acid and its salts;

(-) 3-[4-[2-(3-Oxo-2H-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-ethoxypropanoic acid and its salts;

(±) 3-[6-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]naphthyl]-2-ethoxypropanoic acid and its salts;

(+) 3-[6-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]naphthyl]-2-ethoxypropanoic acid and its salts;

(-) 3-[6-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]naphthyl]-2-ethoxypropanoic acid and its salts;

(±) 3-[6-[2-(2,3-Dihydro-1,4-benzothiazin-4-yl)ethoxy]naphthyl]-2-ethoxypropanoic acid and its salts;

(+) 3-[6-[2-(2,3-Dihydro-1,4-benzothiazin-4-yl)ethoxy]naphthyl]-2-ethoxypropanoic acid and its salts;

(-) 3-[6-[2-(2,3-Dihydro-1,4-benzothiazin-4-yl)ethoxy]naphthyl]-2-ethoxypropanoic acid and its salts;

(±) 3-[4-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-hydroxypropanoic acid and its salts;

(+) 3-[4-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-hydroxypropanoic acid and its salts;

(-) 3-[4-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-hydroxypropanoic acid and its salts;

(±) 3-[4-[2-(2,3-Dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-hydroxypropanoic acid and its salts;

(+) 3-[4-[2-(2,3-Dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-hydroxypropanoic acid and its salts;

(-) 3-[4-[2-(2,3-Dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-hydroxypropanoic acid and its salts;

(±) 3-[4-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-benzyloxypropanoic acid and its salts;

(+) 3-[4-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-

benzyloxypropanoic acid and its salts;

(-) 3-[4-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-benzyloxypropanoic acid and its salts;

(±) 3-[4-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-butoxypropanoic acid and its salts;

(+) 3-[4-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-butoxypropanoic acid and its salts;

(-) 3-[4-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-butoxypropanoic acid and its salts;

(±) 3-[4-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-hexyloxypropanoic acid and its salts;

(+) 3-[4-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-hexyloxypropanoic acid and its salts;

(-) 3-[4-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-hexyloxypropanoic acid and its salts;

(±) 3-[4-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-phenoxypropanoic acid and its salts;

(+) 3-[4-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-phenoxypropanoic acid and its salts;

(-) 3-[4-[2-(2,3-Dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-phenoxypropanoic acid and its salts;

(±) 3-[4-[2-(2,3-Dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-phenoxypropanoic acid and its salts;

(+) 3-[4-[2-(2,3-Dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-phenoxypropanoic acid and its salts;

(-) 3-[4-[2-(2,3-Dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl]-2-phenoxypropanoic acid and its salts;

(±) Methyl 2-methyl-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy] phenyl]-2-phenoxypropanoate;

(+) Methyl 2-methyl-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy] phenyl]-2-phenoxypropanoate;

(-) Methyl 2-methyl-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy] phenyl]-2-phenoxypropanoate;

(±) 2-Methyl-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-phenoxypropanoic acid and its salts;

(+) 2-Methyl-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-phenoxypropanoic acid and its salts;

(-) 2-Methyl-3-[4-[2-(2,3-dihydro-1,4-benzoxazin-4-yl)ethoxy]phenyl]-2-phenoxypropanoic acid and its salts;

(±) Methyl 2-methyl-3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy] phenyl]-2-phenoxypropanoate;

(+) Methyl 2-methyl-3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]

phenyl]-2-phenoxypropanoate;

(-) Methyl 2-methyl-3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy] phenyl]-2-phenoxypropanoate;

(±) 2-Methyl-3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl] -2-phenoxy propanoic acid and its salts;

(+) 2-Methyl-3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl] -2-phenoxy propanoic acid and its salts;

(-) 2-Methyl-3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl] -2-phenoxy propanoic acid and its salts;

(±) 4-Nitrophenyl 3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl] -2-ethoxy propanoate;

(+) 4-Nitrophenyl 3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl] -2-ethoxy propanoate;

(-) 4-Nitrophenyl 3-[4-[2-(2,3-dihydro-1,4-benzothiazin-4-yl)ethoxy]phenyl] -2-ethoxy propanoate;

(±) 3-[4-(4-Benzyl-3,4-dihydro-2H-1,4-benzoxazin-2-yl)methoxyphenyl]-2-ethoxypropanoic acid and its salts;

(+) 3-[4-(4-Benzyl-3,4-dihydro-2H-1,4-benzoxazin-2-yl)methoxyphenyl]-2-ethoxypropanoic acid and its salts;

(-) 3-[4-(4-Benzyl-3,4-dihydro-2H-1,4-benzoxazin-2-yl)methoxyphenyl]-2-ethoxypropanoic acid and its salts;

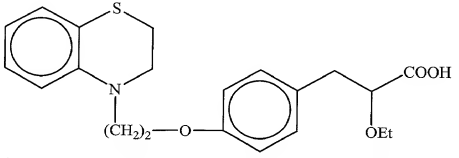
(±) 4-Nitrophenyl-3-[4-(4-benzyl-3,4-dihydro-2H-1,4-benzoxazin-2-yl)methoxy phenyl]-2-ethoxypropanoate;

(+) 4-Nitrophenyl-3-[4-(4-benzyl-3,4-dihydro-2H-1,4-benzoxazin-2-yl)methoxy phenyl]-2-ethoxypropanoate; and

(-) 4-Nitrophenyl-3-[4-(4-benzyl-3,4-dihydro-2H-1,4-benzoxazin-2-yl)methoxy phenyl]-2-ethoxypropanoate; and

a pharmaceutically acceptable carrier, diluent or excipient to a patient in need thereof, wherein the complication is hyperglycemia, osteoporis, hyperlidemia, nephrotic syndrome or disorders related to endothelial cell division.

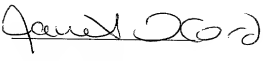
31. A method for preventing or treating complications of diabetes said diabetes caused by insulin resistance or impaired glucose tolerance said method comprising administering a therapeutically effective amount of a 3-[4-[2-(2,3-Dihydro-1, 4-benzothiazin-4-yl)ethoxy] phenyl]-2-ethoxypropanoic acid of the formula:



or a pharmaceutically acceptable salt, polymorph, tautomer, or stereoisomer thereof and a pharmaceutically acceptable carrier, diluent or excipient to a patient in need thereof, wherein the complication is hyperglycemia, osteoporis, hyperlidemia, nephrotic syndrome or

disorders related to endothelial cell division.--

Respectfully submitted,



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